STIC Biotechnology Systems Branch

RAW SEQUENCE LISTING ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number:

Source:

Date Processed by STIC:

er: $\frac{10/578, 493}{1F\omega P}$

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.
PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR CRF SUBMISSION AND PATENTIN SOFTWARE QUESTIONS, PLEASE CONTACT MARK SPENCER, TELEPHONE: 571-272-2510; FAX: 571-273-0221

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE <u>CHECKER</u> <u>VERSION 4.4.0 PROGRAM</u>, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS:

http://www.uspto.gov/web/offices/pac/checker/chkrnote.htm

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail. Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom. Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

- 1. EFS-Bio (http://www.uspto.gov/ebc/efs/downloads/documents.htm, EFS Submission User Manual ePAVE)
- 2. U.S. Postal Service: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450
- Hand Carry, Federal Express, United Parcel Service, or other delivery service (EFFECTIVE 01/14/05):
 U.S. Patent and Trademark Office, Mail Stop Sequence, Customer Window, Randolph Building, 401 Dulany Street,
 Alexandria, VA 22314

Revised 01/10/06

Raw Sequence Listing Error Summary

ERROR DETECTED	SUGGESTED CORRECTION SERIAL NUMBER: 10/578, 493
ATTN: NEW RULES CASES:	PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE
Wrapped Nucleics Wrapped Aminos	The number/text at the end of each line "wrapped" down to the next line. This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3; this will prevent "wrapping."
2Invalid Line Length	The rules require that a line not exceed 72 characters in length. This includes white spaces.
3Misaligned Amino Numbering	The numbering under each 5 th amino acid is misaligned. Do not use tab codes between numbers; use space characters , instead.
4Non-ASCII	The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.
5Variable Length	Sequence(s)contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.
6PatentIn 2.0 "bug"	A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s) Normally, PatentIn would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.
7Skipped Sequences (OLD RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence: (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) (i) SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading) (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) This sequence is intentionally skipped Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to include the skipped sequences.
8Skipped Sequences (NEW RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence. <210> sequence id number <400> sequence id number 000
9Use of n's or Xaa's (NEW RULES)	Use of n's and/or Xaa's have been detected in the Sequence Listing. Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present. In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.
10Invalid <213> Response	Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence. (see item 11 below)
Use of <220>	Sequence(s)missing the <220> "Feature" and associated numeric identifiers and responses. Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section or use "chemically synthesized" as explanation. (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32), also Sec. 1.823 of Sequence Rules
12PatentIn 2.0 "bug"	Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.
13 Misuse of n/Xaa	"n" can only represent a single nucleotide; "Xaa" can only represent a single amino acid



IFWP

RAW SEQUENCE LISTING DATE: 05/17/2006
PATENT APPLICATION: US/10/578,493 TIME: 10:01:13

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

```
3 <110> APPLICANT: Desire, Laurent
      5 <120> TITLE OF INVENTION: BACE455, AN ALTERNATIVE SPLICE VARIANT OF THE HUMAN
             BETA-SECRETASE
     8 <130> FILE REFERENCE: 67987.000002
                                                            Does Not Comply
Corrected Diskette Needed
C--> 10 <140> CURRENT APPLICATION NUMBER: US/10/578,493
C--> 11 <141> CURRENT FILING DATE: 2006-05-05
     13 <160> NUMBER OF SEO ID NOS: 33
    15 <170> SOFTWARE: PatentIn version 3.3
    17 <210> SEQ ID NO: 1
    18 <211> LENGTH: 1368
    19 <212> TYPE: DNA
    20 <213> ORGANISM: Homo sapiens
    22 <400> SEQUENCE: 1
    23 atggcccaag ccctgccctg gctcctgctg tggatggcg cgggagtgct gcctgcccac
                                                                             60
    25 ggcacccage aeggcateeg getgeeeetg egeageggee tggggggege eeeeetgggg
                                                                            120
    180
    29 gtggagatgg tggacaacct gaggggcaag tcggggcagg gctactacgt ggagatgacc
                                                                            240
    31 gtgggcagcc ccccgcagac gctcaacatc ctggtggata caggcagcag taactttgca
                                                                           300
    33 gtgggtgctg ccccccaccc cttcctgcat cgctactacc agaggcagct gtccagcaca
                                                                           360
    35 taccgggacc tccggaaggg tgtgtatgtg ccctacaccc agggcaagtg ggaaggggag
                                                                            420
    37 ctgggcaccg acctggtaag catcccccat ggccccaacg tcactgtgcg tgccaacatt
                                                                           480
    39 gctgccatca ctgaatcaga caagttette atcaaegget ccaaetggga aggeateetg
                                                                           540
    41 gggctggcct atgctgagat tgccaggatc attggaggta tcgaccactc gctgtacaca
                                                                           600
    43 ggcagtctct ggtatacacc catccggcgg gagtggtatt atgaggtcat cattgtgcgg
                                                                           660
    45 gtggagatca atggacagga tctgaaaatg gactgcaagg agtacaacta tgacaagagc
                                                                           720
    47 attgtggaca gtggcaccac caaccttcgt ttgcccaaga aagtgtttga agctgcagtc
                                                                           780
    49 aaatccatca aggcagcctc ctccacggag aagttccctg atggtttctg gctaggagag
                                                                           840
    51 cagetggtgt getggeaage aggeaceace cettggaaca tttteceagt cateteacte
                                                                           900
    53 tacctaatgg gtgaggttac caaccagtcc ttccgcatca ccatccttcc gcagcaatac
                                                                           960
    55 ctgcggccag tggaagatgt ggccacgtcc caagacgact gttacaagtt tgccatctca
                                                                          1020
    57 cagtcatcca cgggcactgt tatgggagct gttatcatgg agggcttcta cgttgtcttt
                                                                          1080
    59 gategggeee gaaaacgaat tggetttget gteagegett geeatgtgea egatgagtte
                                                                          1140
    61 aggacggcag cggtggaagg cccttttgtc accttggaca tggaagactg tggctacaac
                                                                          1200
    63 attecacaga cagatgagte aacceteatg accatageet atgteatgge tgeeatetge
                                                                          1260
    65 geoctettea tgetgecaet etgeeteatg gtgtgteagt ggegetgeet eegetgeetg
                                                                          1320
    67 cgccagcagc atgatgactt tgctgatgac atctccctgc tgaagtga
                                                                          1368
    70 <210> SEQ ID NO: 2
    71 <211> LENGTH: 455
    72 <212> TYPE: PRT
    73 <213> ORGANISM: Homo sapiens
    75 <400> SEQUENCE: 2
    77 Met Ala Gln Ala Leu Pro Trp Leu Leu Trp Met Gly Ala Gly Val
```

5

78 1

RAW SEQUENCE LISTING DATE: 05/17/2006
PATENT APPLICATION: US/10/578,493 TIME: 10:01:13

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

81 Leu Pro Ala His Gly Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser 85 Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp 89 Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val 93 Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr 70 97 Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser 101 Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr 100 105 105 Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val 120 109 Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp 135 113 Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile 150 155 117 Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp 165 170 121 Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Ile Ile Gly 180 185 125 Gly Ile Asp His Ser Leu Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile 195 200 129 Arg Arg Glu Trp Tyr Tyr Glu Val Ile Ile Val Arg Val Glu Ile Asn 130 210 215 133 Gly Gln Asp Leu Lys Met Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser 230 235 137 Ile Val Asp Ser Gly Thr Thr Asn Leu Arg Leu Pro Lys Lys Val Phe 250 245 141 Glu Ala Ala Val Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe 145 Pro Asp Gly Phe Trp Leu Gly Glu Gln Leu Val Cys Trp Gln Ala Gly 275 · 280 149 Thr Thr Pro Trp Asn Ile Phe Pro Val Ile Ser Leu Tyr Leu Met Gly 295 153 Glu Val Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr 310 315 157 Leu Arg Pro Val Glu Asp Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys 325 330 161 Phe Ala Ile Ser Gln Ser Ser Thr Gly Thr Val Met Gly Ala Val Ile 345 340 165 Met Glu Gly Phe Tyr Val Val Phe Asp Arg Ala Arg Lys Arg Ile Gly 360 169 Phe Ala Val Ser Ala Cys His Val His Asp Glu Phe Arg Thr Ala Ala 375 173 Val Glu Gly Pro Phe Val Thr Leu Asp Met Glu Asp Cys Gly Tyr Asn 390 395 177 Ile Pro Gln Thr Asp Glu Ser Thr Leu Met Thr Ile Ala Tyr Val Met

DATE: 05/17/2006

TIME: 10:01:13

```
Input Set : A:\67987.000002.ST25.txt
                  Output Set: N:\CRF4\05172006\J578493.raw
178
                      405
                                             410
181 Ala Ala Ile Cys Ala Leu Phe Met Leu Pro Leu Cys Leu Met Val Cys
                  420
                                        425
185 Gln Trp Arg Cys Leu Arg Cys Leu Arg Gln Gln His Asp Asp Phe Ala
186
           435
                                    440
189 Asp Asp Ile Ser Leu Leu Lys
                                                               Insufficient Emplanation.

Insufficient Emplanation.

What is the Source
What is the Source
What is the Source
I genetic Malenat.

Of genetic Malenat.

Of See Jens II

Sheet.

Sheet.
        450
190
193 <210> SEQ ID NO: 3
194 <211> LENGTH: 6
195 <212> TYPE: PRT
196 <213> ORGANISM: artificial
198 <220> FEATURE:
199 <223> OTHER INFORMATION distinctive fragment
201 <400> SEQUENCE: 3
203 Ile Ala Arg Ile Ile Gly
204 1
207 <210> SEQ ID NO: 4
208 <211> LENGTH: 7
209 <212> TYPE: PRT
210 <213> ORGANISM: artificial
212 <220> FEATURE:
213 <223> OTHER INFORMATION: (distinctive fragment
215 <400> SEQUENCE: 4
217 Glu Ile Ala Arg Ile Ile Gly
218 1
221 <210> SEQ ID NO: 5
222 <211> LENGTH: 8
223 <212> TYPE: PRT
224 <213> ORGANISM: artificial
226 <220> FEATURE:
227 <223> OTHER INFORMATION: distinctive fragment
229 <400> SEQUENCE: 5
231 Glu Ile Ala Arg Ile Ile Gly Gly
232 1
235 <210> SEQ ID NO: 6
236 <211> LENGTH: 8
237 <212> TYPE: PRT
238 <213> ORGANISM: artificial
240 <220> FEATURE:
241 <223> OTHER INFORMATION: distinctive fragment
243 <400> SEQUENCE: 6
245 Ala Glu Ile Ala Arg Ile Ile Gly
246 1
249 <210> SEQ ID NO: 7
250 <211> LENGTH: 9
251 <212> TYPE: PRT
252 <213> ORGANISM: artificial
254 <220> FEATURE:
255 <223> OTHER INFORMATION ✓ AEIARIIGG
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RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/578,493

RAW SEQUENCE LISTING DATE: 05/17/2006 PATENT APPLICATION: US/10/578,493 TIME: 10:01:13

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

```
257 <400> SEQUENCE: 7
259 Ala Glu Ile Ala Arg Ile Ile Gly Gly
260 1
263 <210> SEQ ID NO: 8
264 <211> LENGTH: 10
265 <212> TYPE: PRT
266 <213> ORGANISM: artificial
268 <220> FEATURE:
269 <223> OTHER INFORMATION: distinctive fragment
271 <400> SEQUENCE: 8
273 Ala Glu Ile Ala Arg Ile Ile Gly Gly Ile
274 1
277 <210> SEQ ID NO: 9
278 <211> LENGTH: 9
279 <212> TYPE: PRT
280 <213> ORGANISM: artificial
282 <220> FEATURE:
                                                              Jame Evrol
283 <223> OTHER INFORMATION ( distinctive fragment
285 <400> SEQUENCE: 9
287 Tyr Ala Glu Ile Ala Arg Ile Ile Gly
288 1
291 <210> SEQ ID NO: 10
292 <211> LENGTH: 10
293 <212> TYPE: PRT
294 <213> ORGANISM: artificial
296 <220> FEATURE:
297 <223> OTHER INFORMATION:
                             distinctive fragment
299 <400> SEQUENCE: 10
301 Tyr Ala Glu Ile Ala Arg Ile Ile Gly Gly
302 1
                                         10
305 <210> SEQ ID NO: 11
306 <211> LENGTH: 11
307 <212> TYPE: PRT
308 <213> ORGANISM: artificial
310 <220> FEATURE:
311 <223> OTHER INFORMATION: distinctive fragment
313 <400> SEQUENCE: 11
315 Tyr Ala Glu Ile Ala Arg Ile Ile Gly Gly Ile
316 1
                                         10
319 <210> SEQ ID NO: 12
320 <211> LENGTH: 18
321 <212> TYPE: DNA
322 <213> ORGANISM: artificial
324 <220> FEATURE:
325 <223> OTHER INFORMATION: probe
327 <400> SEQUENCE: 12
328 attgccagga tcattgga
                                                                            18
331 <210> SEQ ID NO: 13
332 <211> LENGTH: 10
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RAW SEQUENCE LISTING DATE: 05/17/2006
PATENT APPLICATION: US/10/578,493 TIME: 10:01:13

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

333	<212> TYPE: DNA	
334	<213> ORGANISM: artificial	
336	<220> FEATURE:	
337	<223> OTHER INFORMATION: primer	
339	<400> SEQUENCE: 13	
340	aggcatcctg	10
	<210> SEQ ID NO: 14	
344	<211> LENGTH: 10	
345	<212> TYPE: DNA	
346	<213> ORGANISM: artificial	
348	<220> FEATURE:	
349	<223> OTHER INFORMATION: primer	
	<400> SEQUENCE: 14	
352	gggctggcct	10
	<210> SEQ ID NO: 15	
	<211> LENGTH: 10	
357	<212> TYPE: DNA	
358	<213> ORGANISM: artificial	
360	<220> FEATURE:	
361	<223> OTHER INFORMATION: primer	
	<400> SEQUENCE: 15	
	atgctgagat	10
	<210> SEQ ID NO: 16	
368	<211> LENGTH: 6	
369	<212> TYPE: DNA	
370	<213> ORGANISM: artificial	
	<220> FEATURE:	
373	<223> OTHER INFORMATION: primer	
	<400> SEQUENCE: 16	
376	tgccag	6
379	<210> SEQ ID NO: 17	
	<211> LENGTH: 6	
381	<212> TYPE: DNA	
382	<213> ORGANISM: artificial	
384	<220> FEATURE:	
385	<223> OTHER INFORMATION: primer	
387	<400> SEQUENCE: 17	
388	gatcat	6
391	<210> SEQ ID NO: 18	
392	<211> LENGTH: 10	
393	<212> TYPE: DNA	
394	<213> ORGANISM: artificial	
396	<220> FEATURE:	
397	<223> OTHER INFORMATION: primer	
	<400> SEQUENCE: 18	
400	tggaggtatc	10
403	<210> SEQ ID NO: 19	
404	<211> LENGTH: 10	
405	<212> TYPE: DNA	

RAW SEQUENCE LISTING ERROR SUMMARY
PATENT APPLICATION: US/10/578,493

DATE: 05/17/2006 TIME: 10:01:14

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

Seq#:33; Xaa Pos. 5

Invalid <213> Response:

Use of "Artificial" only as "<213> Organism" response is incomplete, per 1.823(b) of New Sequence Rules. Valid response is Artificial Sequence.

Seq#:3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22,23,24,25,26,27,28,29 Seq#:30,31,32,33 VERIFICATION SUMMARY

DATE: 05/17/2006

PATENT APPLICATION: US/10/578,493

TIME: 10:01:14

Input Set : A:\67987.000002.ST25.txt
Output Set: N:\CRF4\05172006\J578493.raw

L:10 M:270 C: Current Application Number differs, Replaced Current Application Number

L:11 M:271 C: Current Filing Date differs, Replaced Current Filing Date

L:589 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:33 after pos.:0